Amendments to the Claims:

This listing of claims will replace all prior versions, and listings, of claims in the application:

- 1. (Currently Amended) A pharmaceutical composition comprising a pharmaceutically acceptable carrier and, as active ingredients, an opioid analgesic and a therapeutically effective amount of a compound selected from the group consisting of according to Formula (I)
- (+)-(B)-trans-4-[1-[3,5-bis(trifluoromethyl)benzoyl]-2-(phenylmethyl)-4-piperidinyl]-N-(2,6-dimethylphenyl)-1-piperazine acetamide; and

$$\begin{array}{c|c} R^1 \\ \hline \\ R^2 - X \end{array} \begin{array}{c} R^1 \\ \hline \\ (CH_2)_m \\ (CH_2)_n \end{array} \begin{array}{c} N - L \\ (CH_2)_p \end{array}$$

the pharmaceutically acceptable acid or base addition salts thereof, the stereochemically isomeric forms thereof, the *N*-oxide form thereof<u>.</u> and the prodrugs thereof, wherein

n is 0, 1 or 2;

m is 1 or 2, provided that if m is 2, then n is 1;

p is 1 or 2;

=0 is =0 or $=NR^3$:

X is a covalent bond or a bivalent radical of formula O, S, NR³;

R¹— is Ar¹, Ar¹C₁₋₆alkyl or di(Ar¹)C₁₋₆alkyl, wherein each C₁₋₆alkyl group is optionally substituted with hydroxy, C₁₋₄alkyloxy, oxo or a ketalized oxo substituent of formula O CH₂-CH₂-O or O CH₂-CH₂-CH₂-O;

R² is Ar², Ar²C₁ 6alkyl, Het¹ or Het¹C₁ 6alkyl;

R³ is hydrogen or C₁ 6alkyl;

L is hydrogen; Ar³; C₁ 6alkyl; C₁ 6alkyl substituted with 1 or 2 substituents selected from hydroxy, C₁ 6alkyloxy, Ar³, Ar³C₁ 6alkyloxy and Het²; C₃ 6alkenyl; Ar³C₃ 6alkenyl or a radical of formula

$$-(CHR^{4})_{q}-NR^{5}-C-R^{6}$$

$$-(CHR^{4})_{r}-C-Y^{1}-R^{7}$$

$$-(CHR^{4})_{r}-C-Y^{1}-R^{7}$$
(a-2);
$$R^{8}$$

$$N$$
(a-3);

$$-Y^{2} \xrightarrow{N} A B$$
(a-4); or

$$-(CHR^4)_q - N N - R^3$$
 (a-5);

wherein each q indepen

independently is 2, 3 or 4;

each r is 0, 1, 2, 3 or 4;

each Y¹ independently is a covalent bond, O or NR³;

Y² is a covalent bond, C₁ 4alkanediyl or -C₁ 4alkylNR³-;

each A=B independently is a bivalent radical of formula CH=CH , N=CH or CH=N ;

each R⁴ independently is hydrogen, C₁₋₆alkyl, Ar² or Ar²C₁₋₆alkyl;

R⁵ is hydrogen, C₁ 6alkyl or Ar³;

R⁶ is C₁ 6alkyl, Ar³, Ar³C₁ 6alkyl, di(Ar³)C₁ 6alkyl, Ar³C₃ 7cycloalkyl, or indolyl;

R⁷ is Ar³; Ar³C₁ 6alkyl; di(Ar³)C₁ 6alkyl; C₁ 6alkyl; C₃ 7cycloalkyl; C₃ 7cycloalkyl substituted with Ar³; oxazolyl; oxazolyl substituted with halo

or C1_6alkyl; thiazolyl; thiazolyl substituted with halo or C1_6alkyl; imidazolyl; imidazolyl substituted with Ar³, C1_6alkyl, Ar³C1_6alkyl or halo; indolinyl; indolinyl substituted with C₁ 4alkyl; 2,3,4-trihydroguinolinyl; pyrrolidinyl or furanyl; each R8 independently is hydrogen, C₁₋₆alkyl, C₃₋₇eycloalkyl or a radical of formula of -Alk-R¹¹ (b-1) or $Alk \ Z \ R^{12}$ (b 2): wherein is C_{1 6}alkanedivl: Alk is a bivalent radical of formula -O-, -S- or -NR³: 7___ R-11___ is phenyl; phenyl substituted with 1 or 2 substituents selected from halo. C₁₋₆alkyl or C₁₋₆alkyloxy; furanyl; furanyl substituted with 1 or 2 substituents selected from C₁₋₆alkyl or hydroxyC₁₋₆alkyl; thienyl; thienyl substituted with 1 or 2 substituents selected from halo or C1 6alkyl; oxazolyl; oxazolyl substituted with 1 or 2 C₁₋₆alkyl substituents; thiazolyl; thiazolyl substituted with 1 or 2 C₁ Galkyl substituents; pyridinyl or pyridinyl substituted with 1 or 2 C₁₋₆alkyl substituents; R12 is C₁ 6alkyl or C₁ 6alkyl substituted with hydroxy, carboxyl or C₁₋₆alkyloxycarbonyl; Ar^{1} is phenyl; phenyl substituted with 1, 2 or 3 substituents each independently selected from the group consisting of halo, C1_4alkyl, haloC1_4alkyl, cyano, aminocarbonyl, C1_4alkyloxy and haloC1_4alkyloxy; Ar²is naphtalenyl; phenyl; phenyl substituted with 1, 2 or 3 substituents each independently selected from the group consisting of hydroxy, halo, cyano, nitro, amino, mono- or di(C1 4alkyl)amino, C1 4alkyl, haloC1 4alkyl, C1_4alkyloxy, haloC1_4alkyloxy, carboxyl, C1_4alkyloxycarbonyl, aminocarbonyl and mono- and di(C1 4alkyl)aminocarbonyl; Ar3 is phenyl or phenyl substituted with 1, 2 or 3 substituents selected from the group consisting of halo, hydroxy, amino, nitro, aminocarbonyl, C₁₋₆alkyl, haloC₁ 6alkyl and C₁ 6alkyloxy; Het1. is a monocyclic heterocycle selected from pyrrolyl, pyrazolyl, imidazolyl,

furanyl, thienyl, oxazolyl, isoxazolyl, thiazolyl, isothiazolyl, pyridinyl,

pyrimidinyl, pyrazinyl and pyridazinyl; or a bicyclic heterocycle selected from

the group consisting of quinolinyl, quinoxalinyl, indolyl, benzimidazolyl, benzoxazolyl, benzisoxazolyl, benzothiazolyl, benzisothiazolyl, benzofuranyl and benzothienyl; each monocyclic and bicyclic heterocycle may optionally be substituted on a carbon atom by 1 or 2 substituents selected from the group consisting of halo, C₁ 4alkyl or mono-, di- and tri(halo)methyl; and is a heterocycle selected from the group consisting of 1,4 dihydro-5-oxotetrazol-1-yl, imidazo[1,2-a]pyridinyl, oxazolyl and imidazolyl; each of said

- Het² is a heterocycle selected from the group consisting of 1,4-dihydro-5-oxce tetrazol 1 yl, imidazo[1,2 a]pyridinyl, oxazolyl and imidazolyl; each of heterocycles may be substituted with 1 or where possible 2 substituents selected from the group consisting of C₁-4alkyl and Ar³.
- 2. (Canceled)
- 3. (Canceled)
- 4. (Canceled)
- 5. (Canceled)
- 6. (Canceled)
- 7. (Canceled)
- 8. (Canceled)
- 9. (Withdrawn) A pharmaceutical composition according to claim 1 wherein, the pharmaceutical composition comprises a compound selected from the group consisting of:
- 4-[1-[3,5-bis(trifluoromethyl)benzoyl]-2-(phenylmethyl)-4-piperidinyl]-*N*-(2,6-dimethylphenyl)-1-piperazine acetamide;
- 4-[1-[3,5-bis(trifluoromethyl)benzoyl]-2-(phenylmethyl)-4-piperidinyl]-*N*-(1-phenylcyclohexyl)-1-piperazine acetamide;
- o 1-[3,5-bis(trifluoromethyl)benzoyl]-2-(phenylmethyl)-4-[4-[□-(1-pyrrolidinylcarbonyl)benzyl]-1-piperazinyl]piperidine;

- o 1-[3,5-bis(trifluoromethyl)benzoyl]-4-[4-[1-[(2-methyl-5-oxazolyl)methyl]-1*H*-benzimidazol-2-yl]-1-piperazinyl]-2-(phenylmethyl)piperidine;
- o 4-[1-[3,5-bis(trifluoromethyl)benzoyl]-2-[(4-trifluoromethylphenyl)methyl]-4-piperidinyl]-*N*-(2,6-dimethylphenyl)-1-piperazine acetamide; and
- o 4-[1-[3,5-bis(trifluoromethyl)benzoyl]-2-[(3,4-dichlorophenyl)methyl]-4-piperidinyl]-*N*-(2,6-dimethylphenyl)-1-piperazine acetamide.
- 10. (Currently Amended) A pharmaceutical composition according to claim 1 wherein, the pharmaceutical composition comprises a compound selected from the group consisting of:
- (+) (B) trans-4-[1-[3,5-bis(trifluoromethyl)benzoyl]-2-(phenylmethyl)-4-piperidinyl]-N (2,6-dimethylphenyl) 1-piperazine acetamide;
- (+)-(B)-cis-4-[1-[3,5-bis(trifluoromethyl)benzoyl]-2-(phenylmethyl)-4-piperidinyl]-N-(2,6-dimethylphenyl)-1-piperazine acetamide; and (+)-(B)-trans-4-[1-[3,5-bis(trifluoromethyl)benzoyl]-2-(phenylmethyl)-4-piperidinyl]-N-(2,6-dimethylphenyl)-1-piperazine acetamide (L)-malic acid (1:1).
- 11. (Previously Amended) A pharmaceutical composition according to claim 1 wherein, the pharmaceutical composition is formulated for simultaneous, separate or sequential use.
- 12. (Previously Amended) A pharmaceutical composition according to claim 1 wherein, the opioid analgesic is one or more compounds selected from the group consisting of alfentanil, buprenorphine, butorphanol, carfentanyl, codeine, diacetylmorphine, dihydrocodeine, fentanyl, hydrocodone, hydromorphone, levorphanol, lofentanyl, meperidine, methadone, morphine, nalbuphine, oxycodone, oxymorphone, pentazocine, propoxyphene, remifentanyl and sufentanyl; and derivatives and pharmaceutical acceptable salts thereof.
- 13. (Previously Amended) A pharmaceutical composition according to claim 12 wherein the opioid analysis is one or more compounds selected from the group consisting of oxycodone, codeine, morphine, fentanyl, buprenorphine, hydrocodone, hydromorphone and pharmaceutical acceptable salts and derivatives thereof.

14.	(Previously Amended) A pharmaceutical composition according to claim 1 where, the pharmaceutical composition is in a form suitable to be orally administered.
15.	(Canceled)
16.	(Canceled)
17.	(Canceled)
18.	(Canceled)
19.	(Canceled)
20.	(Canceled)
21.	(Withdrawn) A method for treating pain and/or nociception comprising administering to a person in need thereof an effective amount of a pharmaceutical composition according to claim 1.
22.	(Withdrawn) A method for treating acute and chronic pain selected from the group consisting of inflammatory, post-operative, emergency room (ER), breakthrough, neuropathic and cancer pain comprising administering to one in need thereof an effective amount of a pharmaceutical composition according to claim 1.
23.	(Withdrawn) A method for treating emesis in opioid-based treatments of pain comprising administering to one in need thereof an effective amount of a pharmaceutical composition according to claim 1.
24.	(Withdrawn) A method for treating nausea and vomiting in opioid-based treatments of pain comprising administering to one in need thereof an effective amount of a pharmaceutical composition according to claim 23.
25.	(Withdrawn) A method for treating respiratory depression in opioid-based treatments

of pain comprising administering to one in need thereof an effective amount of an

NK₁-receptor antagonist selected from an NK₁-receptor antagonist according to Formula (I), the pharmaceutically acceptable acid or base addition salts thereof, the stereochemically isomeric forms thereof, the *N*-oxide form thereof and prodrugs thereof.

26. (Withdrawn) A method for reducing and/or overcoming the tolerance observed with opioids in opioid-based treatments of pain comprising administering to one in need thereof an effective amount of an NK₁-receptor antagonist selected from the group consisting of an NK₁-receptor antagonist according to Formula (I), the pharmaceutically acceptable acid or base addition salts thereof, the stereochemically isomeric forms thereof, the *N*-oxide form thereof and prodrugs thereof.